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Attorney Docket No. P67772US1
Application No. 10/509,950**Amendments to the claims:**

This listing of claims replaces all prior versions, and listings, of claims in the application.

Listing of claims:

Claims 1-10 (cancelled).

11 (withdrawn): An antibody specifically immunoreactive with an immunogen, wherein said immunogen is a protein molecule shown in SEQ ID NO. 1, or a fragment, or derivative, or variant thereof.

12 (withdrawn): Use of an antibody of claim 11, for detecting the pathological state of a cell in a sample from a subject, comprising immunocytochemical staining of said cell with said antibody, wherein an altered degree of staining, or an altered staining pattern in said cell compared to a cell representing a known health status indicates a pathological state of said cell.

13 (currently amended): A method of diagnosing ~~or prognosticating~~ Alzheimer's disease in a subject, ~~or determining whether a subject is at increased risk of developing said disease,~~ comprising:

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- determining a ratio level ~~and/or an activity of~~
- (i) a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1), ~~and/or~~
- (ii) a translation product of the gene coding for Htarpp (SEQ ID NO: 1),
- in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus obtained from said subject and
- comparing said ratio level ~~and/or said activity~~ to a reference value representing a known disease or health status, thereby diagnosing ~~or prognosticating~~ Alzheimer's disease in said subject, ~~or determining whether said subject is at increased risk of developing Alzheimer's disease.~~

Claims 14-17 (cancelled).

- 18 (currently amended): The method according to claim 13 wherein said reference value representing a known health status is that of a ratio level ~~and/or an activity of~~
- (i) a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1) ~~and/or~~
- (ii) a translation product of the gene coding for hTARPP (SEQ ID NO: 1),
- in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus sample obtained from a subject not suffering from Alzheimer's disease.

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19 (currently amended): The method according to claim 13 wherein an alteration in the ratio level and/or activity of a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1) and/or a translation product of the gene coding for hTARPP (SEQ ID NO: 1) in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus cell, or tissue, or fluid obtained from said subject relative to a reference value representing a known health status indicates a diagnosis, or prognosis, or increased risk of Alzheimer's disease in said subject.

20 (currently amended): A kit for diagnosing or prognosticating Alzheimer's disease in a subject, or determining the propensity or predisposition of a subject to develop Alzheimer's disease, said kit comprising:

(a) at least one reagent ~~which is~~ selected from the group consisting of

—(i) reagents that selectively detect a transcription product of the gene coding for polypeptide hTARPP (SEQ ID NO: 1), and

—(ii) ~~reagents that selectively detect a translation product of the gene coding for hTARPP (SEQ ID NO: 1), and~~

(b) an instruction for diagnosing, or prognosticating Alzheimer's disease or ~~determining the propensity or predisposition of a subject to develop Alzheimer's disease by~~

— detecting a ratio level, or an activity, or both said level and said activity, of said transcription product and/or said translation product of the gene coding for polypeptide

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hTARPP (SEQ ID NO: 1), in a brain tissue sample of temporal cortex, frontal cortex, and/or hippocampus obtained from said subject[;] and

- ~~diagnosing or prognosticating Alzheimer's disease or determining the propensity or predisposition of said subject to develop Alzheimer's disease, wherein a varied (i) when a ratio level, or activity, or both said level and said activity, of said transcription product is varied and/or said translation product compared to a reference value representing a known health status[,]~~ or wherein a (ii) when a ratio level, or activity, or both said level and said activity, of said transcription product and/or said translation product is similar or equal to a reference value representing a known disease status indicates a diagnosis or prognosis of Alzheimer's disease or an increased propensity or predisposition of developing Alzheimer's disease.

- 21 (withdrawn): A method of treating or preventing a neurodegenerative disease, in particular Alzheimer's disease, in a subject comprising administering to said subject in a therapeutically or prophylactically effective amount an agent or agents which directly or indirectly affect an activity and/or a level of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii).

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- 22 (withdrawn): A modulator of an activity and/or of a level of at least one substance which is selected from the group consisting of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii).
- 23 (withdrawn): Use of a modulator of an activity and/or of a level of at least one substance which is selected from the group consisting of (i) a gene coding for hTARPP, and/or (ii) a transcription product of the gene coding for hTARPP, and/or (iii) a translation product of the gene coding for hTARPP, and/or (iv) a fragment, or derivative, or variant of (i) to (iii) for a preparation of a medicament for treating or preventing a neurodegenerative disease, in particular Alzheimer's disease.
- 24 (withdrawn): A recombinant, non-human animal comprising a non-native gene sequence coding for hTARPP or a fragment, or a derivative, or a variant thereof, said animal being obtainable by:
- (i) providing a gene targeting construct comprising said gene sequence and a selectable marker sequence, and
 - (ii) introducing said targeting construct into a stem cell of a non-human animal, and
 - (iii) introducing said non-human animal stem cell into a non-human embryo, and
 - (iv) transplanting said embryo into a pseudopregnant non-human animal, and
 - (v) allowing said embryo to develop to term, and

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- (vi) identifying a genetically altered non-human animal whose genome comprises a modification of said gene sequence in both alleles, and
- (vii) breeding the genetically altered non-human animal of step (vi) to obtain a genetically altered non-human animal whose genome comprises a modification of said endogenous gene, wherein said disruption results in said non-human animal exhibiting a predisposition to developing symptoms of a neurodegenerative disease or related diseases or disorders.

25 (withdrawn): Use of the recombinant, non-human animal according to claim 24 for screening, testing, and validating compounds, agents, and modulators in the development of diagnostics and therapeutics to treat neurodegenerative diseases, in particular Alzheimer's disease.

26 (withdrawn): An assay for screening for a modulator of neurodegenerative diseases, in particular Alzheimer's disease, or related diseases or disorders of one or more substances selected from the group consisting of

- (i) a gene coding for hTARPP, and/or
- (ii) a transcription product of the gene coding for hTARPP, and/or
- (iii) a translation product of the gene coding for hTARPP, and/or
- (iv) a fragment, or derivative, or variant of (i) to (iii),

said method comprising:

- (a) contacting a cell with a test compound;

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- (b) measuring the activity and/or level of one or more substances recited in (i) to (iv);
- (c) measuring the activity and/or level of one or more substances recited in (i) to (iv) in a control cell not contacted with said test compound; and
- (d) comparing the levels and/or activities of the substance in the cells of step (b) and (c), wherein an alteration in the activity and/or level of substances in the contacted cells indicates that the test compound is a modulator of said diseases or disorders.

Claims 27-31 (cancelled).